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Claims

- 5 1. A pharmaceutical composition comprising a *PRL-1* homologous protein or/and a functional fragment thereof, a nucleic acid molecule encoding a *PRL-1* homologous protein or/and a functional fragment thereof or/and a modulator/effector of said nucleic acid molecule or/and protein together with pharmaceutically acceptable carriers, diluents or/and additives.
- 10 2. The composition of claim 1, wherein the nucleic acid molecule is a vertebrate or insect *PRL-1* nucleic acid, particularly encoding the human *PRL-1* homologs (such as human Prl-1, Prl-2, or Prl-3 protein), and/or a nucleic molecule which is complementary thereto or a functional fragment thereof or a variant thereof.
- 15 3. The composition of claim 1 or 2, wherein said nucleic acid molecule is selected from the group of
- 20 (a) a nucleic acid molecule encoding a polypeptide as shown in Table 2 or/and an isoform, fragment, or/and variant of said polypeptide;
- (b) a nucleic acid molecule which comprised or is the nucleic acid molecule as shown in Table 2;
- (c) a nucleotide sequence which hybridizes at 50°C in a solution containing 1 x SSC and 0.1% SDS to a sequence of (a) or (b),
- 25 (d) a nucleic acid molecule being degenerated as a result of the genetic code to the nucleic acid sequence as defined in (a), (b) or (c);
- (e) a nucleic acid molecule that encodes a polypeptide which is at least 85%, preferably at least 90%, more preferably at least 95%, more preferably at least 98% and up to 99,6% identical to the
- 30 human *PRL-1* homologous protein, preferably as described in Table 2 or as defined in claim 2 or to a polypeptide as defined in (a);
- (f) a nucleic acid molecule that differs from the nucleic acid molecule of (a) to (e) by mutation and wherein said mutation causes an
- 35 alteration, deletion, duplication or premature stop in the encoded

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polypeptide; and

- (g) a partial sequence of any of the nucleotide sequences of (a) to (e) having a length of 15-25 bases, preferably 25-35 bases, more preferably 35-50 bases and most preferably at least 50 bases.

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4. The composition of any one of claims 1-3, wherein the nucleic acid molecule is a DNA molecule, particularly a cDNA or a genomic DNA.

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5. The composition of any one of claims 1-4, wherein said nucleic acid encodes a polypeptide contributing to regulating the energy homeostasis and/or the metabolism of triglycerides.

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6. The composition of any one of claims 1-5, wherein said nucleic acid molecule is a recombinant nucleic acid molecule.

7. The composition of any one of claims 1-6, wherein the nucleic acid molecule is a vector, particularly an expression vector.

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8. The composition of any one of claims 1-5, wherein the polypeptide is a recombinant polypeptide.

9. The composition of claim 8, wherein said recombinant polypeptide is a fusion polypeptide.

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10. The composition of any one of claims 1-7, wherein said nucleic acid molecule is selected from hybridization probes, primers and anti-sense oligonucleotides.

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11. The composition of any one of claims 1-10 which is a diagnostic composition.

12. The composition of any one of claims 1-10 which is a therapeutic composition.

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13. The composition of any one of claims 1-12 for the manufacture of an agent for detecting or/and verifying, for the treatment, alleviation and/or

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prevention of metabolic diseases or dysfunctions, including metabolic syndrome, obesity or/and diabetes, as well as related disorders such as eating disorder, cachexia, hypertension, coronary heart disease, hypercholesterolemia, dyslipidemia, osteoarthritis, gallstones, or liver fibrosis, in cells, cell masses, organs and/or subjects.

14. The composition of any one of claims 1-13 for application in vivo.
15. The composition of any one of claims 1-13 for application in vitro.
16. Use of a nucleic acid molecule encoding a *PRL-1* homologous protein or an isoform, a functional fragment or variant thereof, in particular a nucleic acid molecule as described in Table 2, particularly of a nucleic acid molecule according to claim 3 (a), (b), or (c), or/and a polypeptide encoded thereby or/and a functional fragment or/and a variant of said nucleic acid molecule or said polypeptide or/and a modulator/effector of said nucleic acid molecule or polypeptide for the manufacture of a medicament for the treatment of obesity, diabetes, or/and metabolic syndrome for controlling the function of a gene or/and a gene product which is influenced or/and modified by a *PRL-1* homologous polypeptide.
17. Use of the nucleic acid molecule encoding a *PRL-1* homologous protein or an isoform, a functional fragment or variant thereof, in particular a nucleic acid molecule as described in Table 2, particularly of a nucleic acid molecule according to claim 3 (a), (b), or (c), or/and a polypeptide encoded thereby or/and a functional fragment or/and a variant of said nucleic acid molecule or said polypeptide or/and a modulator/effector of said nucleic acid molecule or said polypeptide for identifying substances capable of interacting with a *PRL-1* homologous polypeptide, particularly with a polypeptide according to claim 3.
18. A non-human transgenic animal exhibiting a modified expression of a *PRL-1* homologous polypeptide, particularly of a polypeptide according to claim 3.

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19. The animal of claim 18, wherein the expression of the *PRL-1* homologous polypeptide, particularly of a polypeptide according to claim 3, is increased and/or reduced.
- 5 20. A recombinant host cell exhibiting a modified expression of a *PRL-1* homologous polypeptide, particularly of a polypeptide according to claim 3.
- 10 21. The cell of claim 20 which is a human cell.
22. A method of identifying a (poly)peptide involved in the regulation of energy homeostasis and/or metabolism of triglycerides in a mammal comprising the steps of
- 15 (a) contacting a collection of (poly)peptides with a *PRL-1* homologous polypeptide, particularly a polypeptide according to claim 3, or a functional fragment thereof under conditions that allow binding of said (poly)peptides;
- (b) removing (poly)peptides which do not bind and
- 20 (c) identifying (poly)peptides that bind to said *PRL-1* homologous polypeptide.
23. A method of screening for an agent which modulates/effects the interaction of a *PRL-1* homologous polypeptide, particularly of a polypeptide according to claim 3, with a binding target/agent, comprising the steps of
- 25 (a) incubating a mixture comprising
- (aa) a *PRL-1* homologous polypeptide, particularly a polypeptide according to claim 3, or a functional fragment thereof;
- 30 (ab) a binding target/agent of said polypeptide or functional fragment thereof; and
- (ac) a candidate agent
- under conditions whereby said polypeptide or functional fragment thereof specifically binds to said binding target/agent
- 35 at a reference affinity;

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- (b) detecting the binding affinity of said polypeptide or functional fragment thereof to said binding target to determine an affinity for the agent; and
- (c) determining a difference between affinity for the agent and the reference affinity.

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24. A method of screening for an agent, which modulates/effects the activity of a *PRL-1* homologous polypeptide, particularly of a polypeptide according to claim 3, comprising the steps of

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- (a) incubating a mixture comprising
 - (aa) said polypeptide or a functional fragment thereof, and
 - (ab) a candidate agentunder conditions whereby said polypeptide or fragment thereof has a reference activity;
- (b) detecting the activity of said polypeptide or functional fragment thereof to determine an activity in presence of the agent; and
- (c) determining a difference between the activity in the presence of the agent and the reference activity.

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20 25. A method of producing a composition comprising the (poly)peptide identified by the method of claim 22 or the agent identified by the method of claim 23 or 24 with a pharmaceutically acceptable carrier, diluent or/and additive.

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26. The method of claim 25 wherein said composition is a pharmaceutical composition for preventing, alleviating or/and treating of metabolic diseases or dysfunctions, including obesity, diabetes, or/and metabolic syndrome, as well as related disorders such as eating disorder, cachexia, hypertension, coronary heart disease, hypercholesterolemia, dyslipidemia, osteoarthritis, gallstones, or liver fibrosis.

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27. Use of a (poly)peptide as identified by the method of claim 22 or of an agent as identified by the method of claim 23 or 24 for the preparation of a pharmaceutical composition for the treatment, alleviation or/and prevention of metabolic diseases or dysfunctions, including obesity, diabetes, or/and metabolic syndrome, as well as related disorders such

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as eating disorder, cachexia, hypertension, coronary heart disease, hypercholesterolemia, dyslipidemia, osteoarthritis, gallstones, or liver fibrosis.

- 5 28. Use of a nucleic acid molecule as defined in any of claims 1-6 or 10 for the preparation of a medicament for the treatment, alleviation or/and prevention of metabolic diseases or dysfunctions, including obesity, diabetes, or/and metabolic syndrome, as well as related disorders such as eating disorder, cachexia, hypertension, coronary heart disease, hypercholesterolemia, dyslipidemia, osteoarthritis, gallstones, or liver fibrosis.
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- 15 29. Use of a polypeptide as defined in any one of claims 1 to 6, 8 or 9 for the preparation of a medicament for the treatment, alleviation or/and prevention of metabolic diseases or dysfunctions, including obesity, diabetes, or/and metabolic syndrome, as well as related disorders such as eating disorder, cachexia, hypertension, coronary heart disease, hypercholesterolemia, dyslipidemia, osteoarthritis, gallstones, or liver fibrosis.
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- 25 30. Use of a vector as defined in claim 7 for the preparation of a medicament for the treatment, alleviation or/and prevention of metabolic diseases or dysfunctions, including obesity, diabetes, or/and metabolic syndrome, as well as related disorders such as eating disorder, cachexia, hypertension, coronary heart disease, hypercholesterolemia, dyslipidemia, osteoarthritis, gallstones, or liver fibrosis.
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31. Use of a host cell as defined in claim 20 or 21 for the preparation of a medicament for the treatment, alleviation or/and prevention of metabolic diseases or dysfunctions, including obesity, diabetes, or/and metabolic syndrome, as well as related disorders such as eating disorder, cachexia, hypertension, coronary heart disease, hypercholesterolemia, dyslipidemia, osteoarthritis, gallstones, or liver fibrosis.
- 35 32. Use of a *PRL-1* homologous nucleic acid molecule or/and of a functional fragment thereof for the production of a non-human transgenic animal which over- or under-expresses the *PRL-1*

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homologous gene product.

33. Kit comprising at least one of

- 5 (a) a *PRL-1* homologous nucleic acid molecule or/and a functional fragment thereof;
- (b) a *PRL-1* homologous amino acid molecule or/and a functional fragment or/and an isoform thereof;
- (c) a vector comprising the nucleic acid of (a);
- (d) a host cell comprising the nucleic acid of (a) or the vector of (c);
- 10 (e) a polypeptide encoded by the nucleic acid of (a);
- (f) a fusion polypeptide encoded by the nucleic acid of (a);
- (g) an antibody, an aptamer or/and another modulator/effector of the nucleic acid of (a) or the polypeptide of (b), (e), or/and (f) and
- (h) an anti-sense oligonucleotide of the nucleic acid of (a).

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